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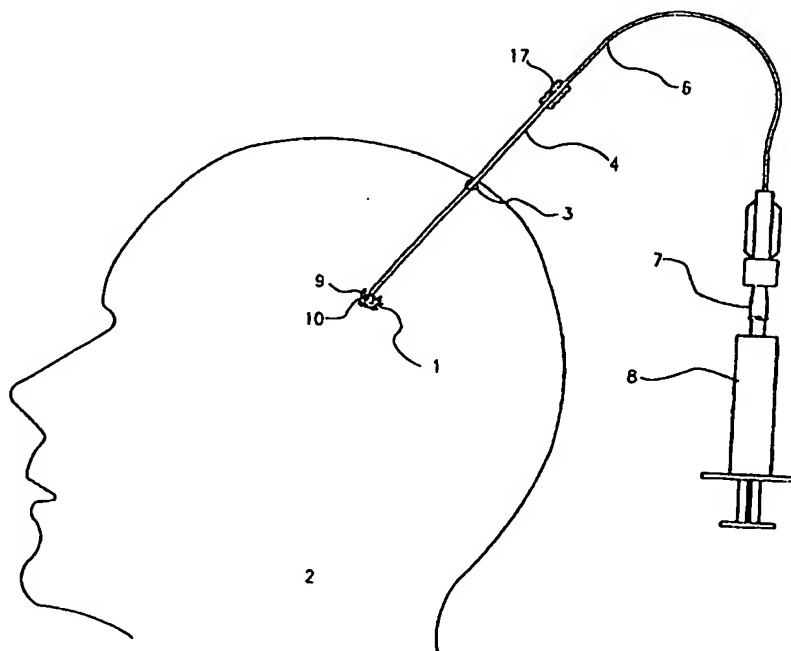
INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

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(21) International Application Number: PCT/SE97/00492 (22) International Filing Date: 21 March 1997 (21.03.97) (30) Priority Data: 9601974-0 23 May 1996 (23.05.96) SE (71) Applicant (for all designated States except US): ELEKTA AB [SE/SE]; P.O. Box 7593, S-103 93 Stockholm (SE). (72) Inventors; and (75) Inventors/Applicants (for US only): HÅKANSSON, Håkan [SE/SE]; Astrakanvägen 6, S-224 56 Lund (SE). OLS-SON, Jan-Erik [SE/SE]; Hammarkindsgatan 8, S-582 47 Linköping (SE). WALLINDER, Jonas [SE/SE]; Smedstorp, Ledberg, S-590 47 Vikingstad (SE). (74) Agent: AWAPATENT AB; P.O. Box 45086, S-104 30 Stockholm (SE).		(81) Designated States: AL, AM, AT, AT (Utility model), AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, CZ (Utility model), DE, DE (Utility model), DK, DK (Utility model), EE, EE (Utility model), ES, FI, FI (Utility model), GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK (Utility model), TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG). Published <i>With international search report.</i>

(54) Title: DEVICE FOR INJECTING A SUBSTANCE INTO A BODY, ESPECIALLY HUMAN OR ANIMAL

(57) Abstract

A device for injecting a substance into a body, especially human or animal, comprising a cannula or needle member (4-6) and a dosing device (8) for supplying a determined amount of substance (10) to a target area (1), the dosing device having an attachment (7) for the cannula or needle member. The invention is characterised in that the cannula or needle member (4-6) comprises a tube (4) and a catheter (6) which is displaceable in the tube, the outer diameter of the catheter being smaller than the inner diameter of the tube, thereby forming a gap between the catheter (5) and the tube (4), essentially over the entire length of the tube (4), in that the proximal portion of the catheter is attached to the dosing device (8), and in that the distal portion (9) of the catheter, in connection with said injection, is arranged adjacent to the distal end (12) of the tube (4) and forms a fluid-tight seal therewith.



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DEVICE FOR INJECTING A SUBSTANCE INTO A BODY, ESPECIALLY HUMAN OR ANIMAL

The present invention relates to a device for injecting a substance into a body, especially human or animal, comprising a cannula or needle member and a dosing device for delivering a determined amount of substance to a target area, the dosing device having an attachment for the cannula or needle member. The invention also concerns a method for injecting the substance, comprising the steps of defining the target area, inserting a cannula or needle member in the body so as to position the tip portion of the member at the target area, supplying a determined amount of substance to the target area, and removing the cannula or needle member.

In the medico-technical field, several methods of introducing liquid, with or without solid particles, into the body are already available. Most of these methods are based on the use of a needle or catheter in combination with some pumping device such as syringes or infusion pumps. In most applications, the volume of infused or injected liquid, liquid mixture or suspension is also relatively great relative to the dimensions or inner volume of the insertion system, which results in the influence of the system on the final position of the inserted fluid volume in the body being affected only marginally when withdrawing the insertion system. In most applications, the position of the deposition is also not decisive of the clinical efficiency of the infused substance.

In recent years, the requirements for accuracy of the deposition have increased dramatically in the field of, among other things, tissue implants. An example of fields where the accuracy of the deposition is highly decisive of the clinical result is the implantation of cells for treating Parkinson's disease. Cells which preferably are extracted from a foetus are suspended in a

physiological solution, which is inserted to the desired position in the brain. The amount of suspended cells is in the order of 10 μ l, which corresponds to the volume in a 10 mm length of the catheter by which the substance is introduced. Problems arise when the catheter system is being withdrawn, since a vacuum is generated which sucks back a great part of the infused amount to a non-desired position, resulting in the fact that the implanted cells cannot develop and contribute to a positive clinical effect.

The object of the present invention is to satisfy the requirements described above, while eliminating the drawbacks of prior-art systems.

A further object is to provide a method and a device, by which a substance which in medical contexts is suitable, can be injected with accuracy and in its entirety into a restricted and well-defined target area in a patient's body.

One more object of the invention is to permit the use of a flexible catheter for a simplified injection process.

According to the invention, these objects are achieved by a device according to the introductory part, which is characterised in that the cannula or needle member comprises a tube and a catheter displaceable in said tube, the outer diameter of the catheter being smaller than the inner diameter of the tube, thereby forming a gap between the catheter and the tube, essentially over the entire length of the tube, in that the proximal portion of the catheter is attached to the dosing device, and in that the distal portion of the catheter is, in connection with said injection, arranged adjacent to the distal end of the tube and forms a fluid-tight seal therewith.

A method for manipulating the device as described above is characterised in that the cannula or needle member is inserted into the body, the tip portion thereof

being sealed by means of a sealing member which forms a fluid-tight seal with a surrounding tube of the cannula or needle member, in that the substance is injected into the target area by means of a catheter, which is surrounded by said tube and whose distal portion has, on the occasion of injection, been brought close to the distal portion of the tube, in that the catheter is then withdrawn from the body, thereby allowing air or gas at essentially atmospheric pressure to enter the space in the tube between the target area and the distal portion of the catheter, and in that finally the tube is withdrawn from the body and the insertion opening for the cannula or needle member is closed.

Further developments of the invention appear from the features stated in the subclaims.

Preferred embodiments of the invention will now be described for the purpose of exemplification, reference being made to the accompanying drawings, in which:

Fig. 1 illustrates an embodiment of the inventive device inserted to the target area, in a patient's brain, but before injecting a medical substance;

Fig. 2 is the same view as in Fig. 1, illustrating a dosing device with a catheter connected, whose distal end is positioned adjacent to the target area, after injecting the substance;

Fig. 3 illustrates on a larger scale the tip portion of the cannula or needle member in the target area during injection; and

Fig. 4 is the same view as in Fig. 3, illustrating an alternative embodiment of the device according to the invention.

With reference first to Figs 1-3, the injection or infusion device according to the invention comprises an outer tube 4 of a suitable surgical material. Preferably, the tube is rigid and made of stainless steel. The distal tube portion 12, which is adapted to be positioned adjacent to a target area 1 in the patient's body, has in

this embodiment a smaller diameter and a considerably shorter length than the proximal tube portion 13, which projects from the patient's body. In the preferred embodiment, the length of the entire tube is about 200 mm and the length of the area having a smaller diameter is about 1 mm. The outer diameter of the tube is about 2 mm and its inner diameter is about 1.6 mm except in the area of a smaller diameter, where the inner diameter is about 1.3 mm. In Fig. 3, the difference in diameter is clearly shown, and the reason for this difference will be explained below.

Thus, the tube 4 is adapted to be inserted into a patient's (a human or an animal) body and in order to facilitate this, the proximal portion of the tube can be fitted with a finger grip portion 17, as illustrated in Fig. 1. This figure shows the tube 4 inserted in the patient's skull 2 for injecting a cell suspension into a target area 1 in the brain as above, but it is to be understood that the inventive device can just as well be used to inject an optional substance which is suitable in medical contexts, such as a pharmaceutical preparation, cytotoxin, contrast fluid etc, into a patient's internal organs, spinal marrow, tissue etc. The main inventive object is injection of a small amount of substance into a well-defined target area.

Reference is again made to Fig. 1, illustrating that the tip portion or distal portion 12 of the tube 4 has been positioned at the target area 1. When inserting the tube 4 into the skull 2, a mandrin 5 is stationarily and releasably arranged in the tube. The mandrin is formed as a cylindrical rod of uniform thickness, preferably of the same material as, but longer than the tube 4 and also having a finger grip portion 18 at its proximal portion. The outer diameter of the mandrin 5 is the same as the inner diameter of the distal tube portion 12, in this case about 1.3 mm. This results in a seal against the inner side of the tube in the distal portion 12 thereof

against penetration of cerebral substance, blood, tissue material and the like into the tube 4 when inserting it into the skull 2. Moreover, an essentially concentric gap 14 is formed between the mandrin 5 and the tube portion having a greater diameter 13, which gap is open to the atmosphere. If desired, the gap can alternatively be connected to a gas source (not shown) having a pressure slightly exceeding atmospheric pressure for reasons that will be explained below. If necessary, an additional sealing member, e.g. an O-ring, can be arranged in prior-art manner on the distal end of the mandrin (not shown).

When the tube 4 has been positioned, the mandrin 5 is withdrawn from the tube, and initially a vacuum arises in the distal portion 12 of the tube, but owing to the small length of this portion and owing to the gap 14 which serves as a pressure-equalising member, the vacuum is eliminated almost immediately, and therefore essentially no matter will be sucked into the tube 4.

The inventive device also comprises a catheter 6, preferably in the form of a plastic tube, whose outer diameter is the same as that of the mandrin, i.e. in this case about 1.3 mm, so as to obtain the same sealing effect as with the mandrin. Its inner diameter (lumen) is in this example about 0.9 mm and its length in the order of 300-400 mm. The proximal portion of the catheter is, for instance by means of a so-called Luer connection 7, attached to a dosing device/injection device 8, which in Fig. 2 is schematically illustrated as a syringe.

After removing the mandrin 5 from the tube 4, the catheter 6 is inserted therein and is guided thereby. The air or gas in the tube flows out through the above-mentioned gap 14, which is now delimited by the inner diameter of the tube 4 and the outer diameter of the catheter (which is the same as that of the mandrin). The catheter 6 is preferably inserted so far that its tip is positioned immediately outside the tip of the tube, as shown in Fig. 3. If necessary, additional sealing means,

for instance ridges 11 as indicated in Fig. 3, can be arranged on the distal portion of the catheter. When the catheter 6 has been positioned, a determined amount of substance 10, e.g. a cell suspension, is infused or injected into the target area 1 by means of the dosing device/injection device 8. When the infusion has been carried out, the catheter 6 is withdrawn from the tube 4, whereby pressure equalisation occurs in the same manner as described above in connection with the withdrawal of the mandrin 5. Then the tube 4 communicating with the atmosphere or a gas source as stated above is withdrawn, which means that there will be no negative pressure.

In Fig. 3, reference numeral 11 indicates a number of parallel ribs or ridges which are circumferentially oriented. In another embodiment, reference numeral 11 indicates, however, an X-ray marker for identifying the position of the catheter end in the body.

Fig. 4 illustrates an alternative embodiment of the device according to the invention. What essentially distinguishes this embodiment from the one previously presented is that it comprises no mandrin. The catheter 6 is in place even when inserting the cannula or needle member, i.e. in this case, the tube 4. The tube can also in this case be of the design presented above - a short distal portion having a small inner diameter and a long proximal portion having a greater inner diameter - but a tube having a constant inner diameter is preferred. At the distal portion of the catheter 6, a piston-shaped member 15 is arranged in a fixed or integrated manner on the outer surface of the catheter. The piston-shaped member 15 is slidingly engaged with the inner surface of the tube 4 and serves as a sealing member for the gap 14 between the tube and the catheter. In its cylindrical peripheral surface, the member 15 has a plurality of essentially axially oriented grooves or channels 16 of such small dimensions that it does not allow the passing of substance or fluid, but allows the passing of air or

gas, thereby preventing any matter from penetrating into the gap 14 when inserting the cannula or needle member, but permitting pressure equalisation when the catheter after the injection is being withdrawn from the tube 4.

5 Penetration of any matter into the lumen of the catheter during said insertion is counteracted by the fluid column therein. The rigidity of the plastic material of the catheter 6 is such as to allow the counterpressure during said insertion to be absorbed by the catheter, and its
10 lateral outward bend, if any, is restricted by the surrounding tube 4.

The method for injecting or infusing a substance into a patient by the inventive device has been described above in broad outline, but a brief description may
15 be justified.

First the size and position of the target area in the body are defined and then the cannula or needle member is inserted into the body so as to position its tip proportion at the target area 1.

20 In the Example illustrated in Figs 1 and 2, this takes place in such a manner that a hole 3 is made in the cranial bone and the tube 4 with the retracted mandrin 5 or catheter 6 and the piston member 15 are positioned in the relevant area in the brain. This is carried
25 out suitably by means of a device as described in ELEKTA INSTRUMENT AB's European Patent Application 312,568 (US Patent 5,116,344) or by means of the above-mentioned marker 11. In the embodiment shown in Fig. 2, the catheter is filled with a solution, whereupon a calculated
30 dose of cell suspension 10 is sucked into the distal portion of the catheter. When the mandrin has been withdrawn from the tube 4, the charged catheter 6 in the tube 4 is inserted in position, cf. Fig. 3, for instance by means of a marking or scale on the outer surface of the catheter
35 on the same level as the proximal end of the tube, cf. Fig. 2. The cell suspension 10 is then infused by means of the injection device 8, and the catheter 6 is

removed from the body (tube) during pressure equalisation as described above. Then the tube 4 is removed and the insertion opening 3 is sealed.

5 In the embodiment shown in Fig. 4, the substance (cell suspension) is already in place in the distal portion of the catheter when the tube 4 along with the catheter 6 is inserted in the patient.

The invention is not restricted to that described above or shown in the drawings, but can be modified with-
10 in the scope of the appended claims.

CLAIMS

1. A device for injecting a substance into a body,
5 especially human or animal, comprising a cannula or
needle member (4-6) and a dosing device (8) for deliver-
ing a determined amount of substance (10) to a target
area (1), the dosing device having an attachment (7) for
the cannula or needle member, c h a r a c t e r i s e d
10 in that the cannula or needle member (4-6) comprises a
tube (4) and a catheter (6) displaceable in said tube,
the outer diameter of the catheter being smaller than
the inner diameter of the tube, thereby forming a gap
(14) between the catheter (6) and the tube (4), essen-
15 tially over the entire length of the tube (4), in that
the proximal portion of the catheter is attached to the
dosing device (8), and in that the distal portion (9)
of the catheter is, in connection with said injection,
arranged adjacent to the distal end (12) of the tube (4)
20 and forms a fluid-tight seal therewith.
2. The device as claimed in claim 1, c h a r a c -
t e r i s e d in that the gap (14) is open to the atmos-
phere or connected to a gas source having a low positive
pressure.
- 25 3. The device as claimed in claim 1 or 2, c h a r -
a c t e r i s e d in that a sealing member (11, 15) is
arranged on the distal portion (9) of the catheter (6) in
sealing engagement with the interior of the tube (4).
- 30 4. The device as claimed in any one of the preceding
claims, c h a r a c t e r i s e d in that the tube (4)
has a short distal portion (12) having a smaller inner
diameter than the remainder (13) of the tube, and in that
said inner diameter is essentially as large as the outer
diameter of the catheter (6).
- 35 5. The device as claimed in claim 1, c h a r a c -
t e r i s e d in that the cannula or needle member (4-6)
also comprises a mandrin (5) displaceable in the tube (4)

and being in the form of a cylindrical rod having essentially the same outer diameter as the catheter (6) and a greater length than the tube (4), in that the mandrin (5) is arranged to exchangeably replace the catheter (6) when
5 inserting the cannula or the needle member (4-6) in said body, and in that one end of the mandrin (5) is arranged adjacent to the distal portion (12) of the tube (4) and forms a fluid-tight seal therewith in connection with said insertion.

10 6. The device as claimed in any one of claims 1-4, characterised in that the sealing member (11, 15) is a piston-shaped member (15) which is arranged, in a fixed or integrated manner, on the outside of the distal portion of the catheter (6) and which has small,
15 essentially axially directed grooves dimensioned so as not to allow fluid to pass, but to allow gas to pass.

7. The device as claimed in any one of the preceding claims, characterised in that the catheter (6) is made of a flexible tube material.

20 8. A method for injecting a substance into a body, especially human or animal, comprising the steps of defining the target area (1), introducing a cannula or needle member (4-6) into the body so as to position the tip portion (9-12) of the member (4-6) at the target
25 area, delivering a determined amount of substance (10) to the target area, and removing the cannula or needle member (4-6), characterised in that the cannula or needle member (4-6) is inserted into the body, the tip portion (9, 12) thereof being sealed by means of
30 a sealing member (5, 15) which forms a fluid-tight seal with a surrounding tube (4) of the cannula or needle member (4-6), in that the substance is injected into the target area (1) by means of a catheter (6), which is surrounded by said tube (4) and whose distal portion (9)
35 has, on the occasion of injection, been brought close to the distal portion (12) of the tube (4), in that the catheter (6) is then removed from the body, thereby

allowing air or gas at essentially atmospheric pressure to enter the space in the tube (4) between the target area (1) and the distal portion (9) of the catheter (6), and in that finally the tube (4) is removed from the body and the insertion opening (3) for the cannula or needle member (4-6) is closed.

9. The method as claimed in claim 8, c h a r a c - t e r i s e d in that during said insertion, the cannula or needle member (4-6) comprises a sealing mandrin (5) of the same outer diameter as the catheter (6), and in that after said insertion, the mandrin (5) is replaced by the catheter (6) for injection.

10. The method as claimed in claim 8, c h a r a c - t e r i s e d in that during said insertion, the cannula or needle member (4-6) comprises the catheter (6) which at its distal portion (9) is provided with a piston-shaped element (15) in sealing engagement with the surrounding tube (4) and which is then stationary relative to the tube (4), and in that injection takes place through the still stationary catheter (6).

1/4

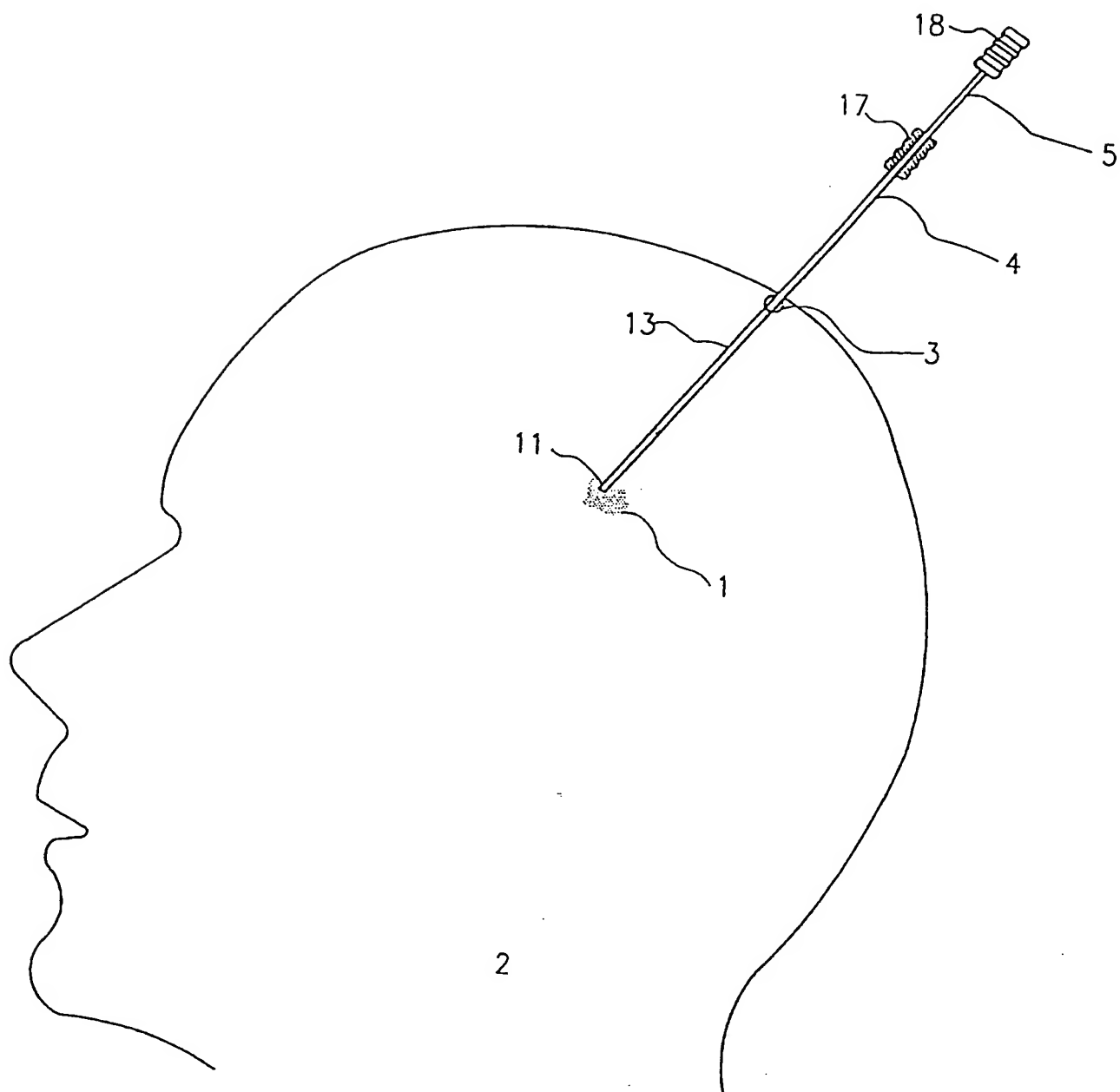


Fig. 1

SUBSTITUTE SHEET

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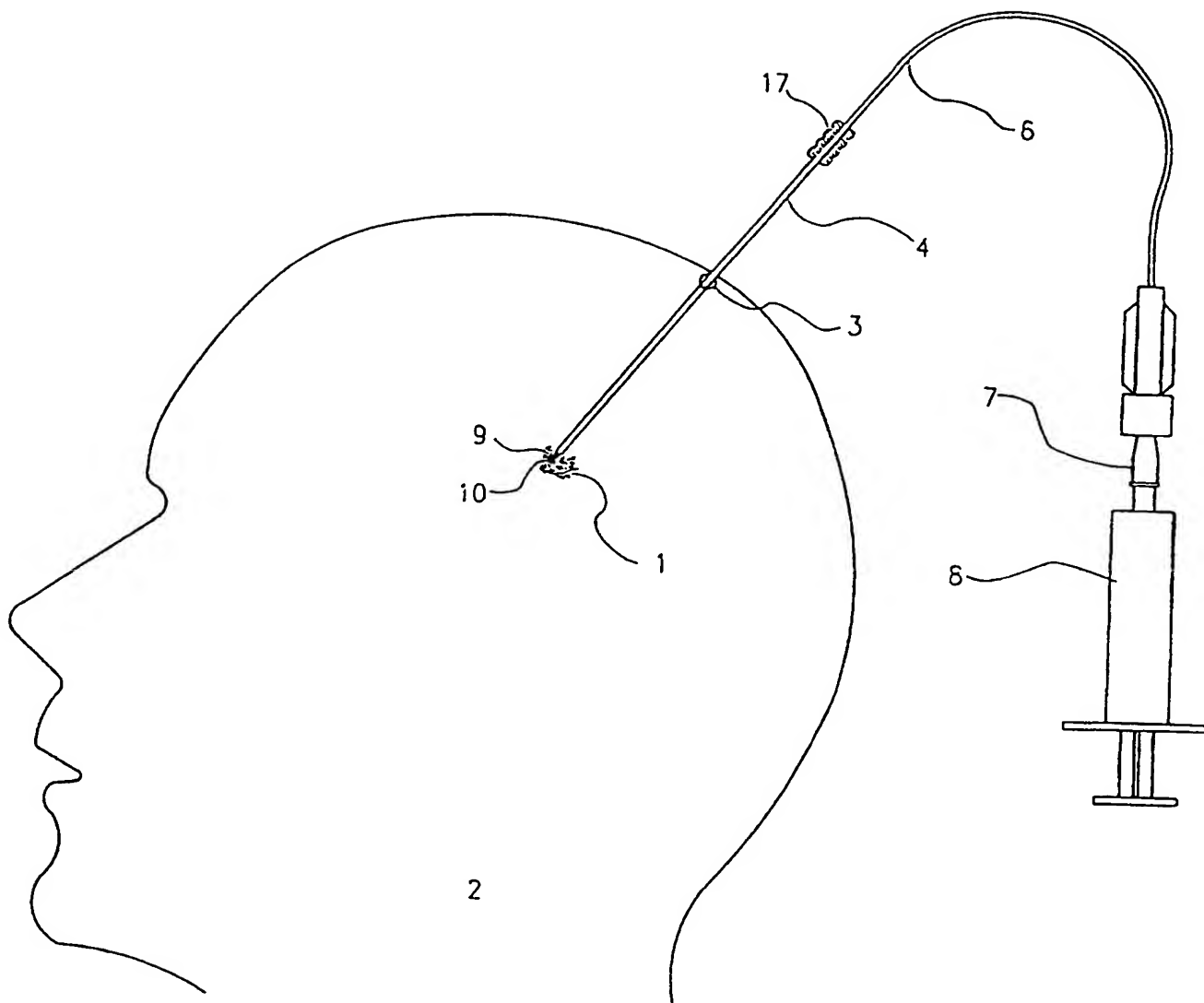


Fig. 2

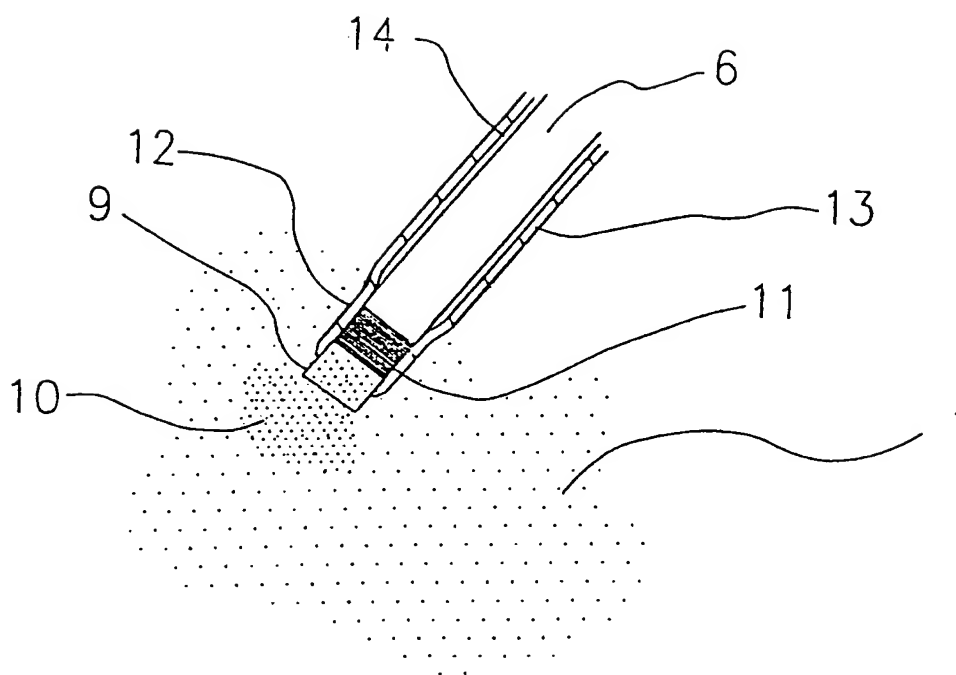


Fig. 3

4/4

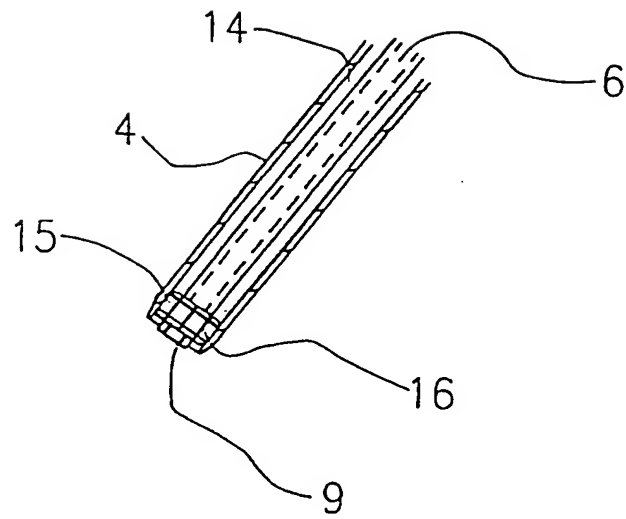


Fig. 4

INTERNATIONAL SEARCH REPORT

International application No.

PCT/SE 97/00492

A. CLASSIFICATION OF SUBJECT MATTER

IPC6: A61M 5/32, A61M 5/158

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC6: A61M

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched
SE,DK,FI,NO classes as above

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

DIALOG: WPI, CLAIMS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	DE 3539243 C3 (OMNI MEDICAL LTD), 12 December 1991 (12.12.91), column 4, line 14 - line 38, figure 1 -- -----	1-7

☐ Further documents are listed in the continuation of Box C.☒ See patent family annex.

* Special categories of cited documents:

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "B" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

- "X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

- "Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

- "&" document member of the same patent family

Date of the actual completion of the international search

23 June 1997

Date of mailing of the international search report

05 -07- 1997

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INTERNATIONAL SEARCH REPORT

International application No.

PCT/SE 97/00492**Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)**

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.: **8-10**
because they relate to subject matter not required to be searched by this Authority, namely:

Method for treatment of the human or animal body by surgery or therapy PCT Rule 67.1(iv)
2. ☐ Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest☐

The additional search fees were accompanied by the applicant's protest.

☐

No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT
Information on patent family members

03/06/97

International application No.
PCT/SE 97/00492

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
DE 3539243 C3	12/12/91	NONE	